

# Efficient estimation of modified treatment policy effects based on the generalized propensity score

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## Motivating example

The *observed data* unit is  $O := (W, A, Y) \sim P_0 \in \mathcal{M}$ :

- $W \in \mathbb{R}^d$  is a vector of baseline covariates;
- $A \in \mathbb{R}$  is a continuous-valued exposure; and
- $Y \in \mathbb{R}$  is an outcome of interest.

Let  $\mathcal{M}$  be a large *semiparametric model* and for each  $P \in \mathcal{M}$ , define the *population intervention effect* (PIE) as

$$\Psi_\delta(P) := \mathbb{E}_P\{Y(A_\delta) - Y\},$$

where  $A_\delta$  arises from a *stochastic* intervention.

## NPSEM with static interventions

- Use a nonparametric structural equation model (NPSEM) to describe the generation of  $O$  (Pearl 2009), specifically

$$W = f_W(U_W); A = f_A(W, U_A); Y = f_Y(A, W, U_Y)$$

- Implies a model for the distribution of counterfactual random variables generated by interventions on the process.
- A *static intervention* replaces  $f_A$  with a specific value  $a$  in its conditional support  $A | W$ .
- This requires specifying a particular value of the exposure under which to evaluate the outcome *a priori*.

## NPSEM with stochastic interventions

- *Stochastic interventions* modify the value  $A$  would naturally assume by drawing from a modified exposure distribution.
- Consider the post-intervention value  $A_\delta \sim G_\delta(\cdot | W)$ ; static interventions are a special case (degenerate distribution).
- Such an intervention generates a counterfactual RV  $Y_{G_\delta} := f_Y(A_\delta, W, U_Y)$ , with distribution  $P_0^\delta$ .
- We aim to estimate  $\psi_{0,\delta} := \mathbb{E}_{P_0^\delta}\{Y_{G_\delta}\}$ , the counterfactual mean under the post-intervention exposure distribution  $G_\delta$ .

## Stochastic interventions for the causal effects of shifts

- Díaz and van der Laan (2012; 2018)'s *stochastic* interventions

$$\delta(a, w) = \begin{cases} a + \delta, & a + \delta < u(w) \quad (\text{if plausible}) \\ a, & a + \delta \geq u(w) \quad (\text{otherwise}) \end{cases}$$

- Haneuse and Rotnitzky (2013): *modified treatment policies*

- Evaluate outcome under modified *intervention distribution*:

$$P^\delta(g_{0,A})(A = a \mid W) = g_{0,A}(\delta^{-1}(A, W) \mid W).$$

- Díaz and van der Laan (2018) show that  $\psi_{0,\delta}$  is identified by a functional of the distribution of  $O$ :

$$\psi_{0,\delta} = \int_{\mathcal{W}} \int_{\mathcal{A}} \mathbb{E}_{P_0}\{Y \mid A = \delta(a, w), W = w\} \cdot g_{0,A}(a \mid W = w) \cdot q_{0,W}(w) d\mu(a) d\nu(w)$$

## Estimation of the PIE

An estimator  $\psi_n$  of  $\psi_0 := \Psi(P_0)$  is *efficient* if and only if

$$\psi_n - \psi_0 = n^{-1} \sum_{i=1}^n D^*(P_0)(O_i) + o_P(n^{-1/2}),$$

where  $D^*(P)$  is the *efficient influence function* (EIF) of  $\Psi_\delta$  with respect to the model  $\mathcal{M}$  at  $P$ .

The EIF of  $\Psi$  is indexed by two key *nuisance parameters*

$$\bar{Q}_{P,Y}(A, W) := \mathbb{E}_P(Y | A, W) \quad \text{outcome mechanism}$$

$$g_{P,A}(A, W) := p(A | W) \quad \text{generalized propensity score}$$

## Estimation of a counterfactual mean

We'll rely on *empirical process notation* throughout:

- $P_0 f = \mathbb{E}_{P_0} \{f(O)\} = \int f(o) dP(o)$
- $P_n f = \mathbb{E}_{P_n} \{f(O)\} = n^{-1} \sum_{i=1}^n f(O_i)$

We can estimate the *counterfactual mean*  $\Psi_\delta(P)$ , using the inverse probability weighted (IPW) estimator

$$\psi_{\delta,n} = n^{-1} \sum_{i=1}^n \frac{g_{n,A}(\delta^{-1}(A_i, W_i) | W_i)}{g_{n,A}(A_i | W_i)} Y_i.$$

## Why IPW estimators?

- IPW estimators are the oldest class of causal effect estimators.
- IPW estimators are still very commonly used in practice today.
- Easy to implement and appropriate in many settings, but...
  1. requires a correctly specified estimate of the propensity score;
  2. can be inefficient, never attaining the efficiency bound; and
  3. suffers from an (asymptotic) curse of dimensionality.

## IPW estimators

The IPW estimator  $\Psi_\delta(P_n, g_{n,A})$  is a solution to the score equation  $D_{\text{IPW}}(O; \Psi_\delta) = \frac{(g_{n,A}(\delta^{-1}(A_i, W_i) | W_i))}{g_{n,A}(A_i | W_i)} Y - \Psi(P)$ :

$$\Psi_\delta(P_n, g_{n,A}) = n^{-1} \sum_{i=1}^n \frac{g_{n,A}(\delta^{-1}(A_i, W_i) | W_i)}{g_{n,A}(A_i | W_i)} Y_i.$$

- Consistency and convergence rate of IPW relies on those same properties of the generalized propensity score estimator  $g_{n,A}$ .
- Generally, finite-dimensional (i.e., parametric) models are not flexible enough to consistently estimate  $g_{0,A}$ .

## Nonparametric conditional density estimation

- Our IPW estimator require the generalized propensity score, at both  $g_A(A | W)$  and  $g_A(\delta^{-1}(A, W) | W)$ .
- There is a rich literature on density estimation, we follow the approach first explored in Díaz and van der Laan (2011).
- To build a conditional density estimator, consider

$$g_{n,A,\alpha}(A | W) = \frac{\mathbb{P}(A \in [\alpha_{t-1}, \alpha_t] | W)}{|\alpha_t - \alpha_{t-1}|}.$$

- This is a classification problem, where we estimate the probability that a value of  $A$  falls in a bin  $[\alpha_{t-1}, \alpha_t)$ .
- The choice of the tuning parameter  $t$  corresponds roughly to the choice of bandwidth in classical kernel density estimation.

## Nonparametric conditional density estimation

- Díaz and van der Laan (2011) propose a reformulation of this classification approach as a set of hazard regressions.
- To effectively employ this proposed reformulation, consider

$$\mathbb{P}(A \in [\alpha_{t-1}, \alpha_t) \mid W) = \mathbb{P}(A \in [\alpha_{t-1}, \alpha_t) \mid A \geq \alpha_{t-1}, W) \times \prod_{j=1}^{t-1} \{1 - \mathbb{P}(A \in [\alpha_{j-1}, \alpha_j) \mid A \geq \alpha_{j-1}, W)\}$$

- Likelihood may be re-expressed as the likelihood of a binary variable in a repeated measures data structure.
- Specifically, the observation of  $O_i$  is repeated as many times as intervals  $[\alpha_{t-1}, \alpha_t)$  are prior to the interval to which  $A_i$  falls, and the indicator variables  $A_i \in [\alpha_{t-1}, \alpha_t)$  are recorded.

## Curse of dimensionality

Goal: Construct nuisance parameter *estimators* that are *consistent* and *converge faster* than  $n^{-1/4}$  under *minimal assumptions*.

*Challenging* for moderately large  $d$ , i.e., *curse of dimensionality*.

For example, consider *kernel regression* with bandwidth  $h$  and kernels orthogonal to polynomials in  $W$  of degree  $k$ .

- Assume parameter is  $k$  times *differentiable*.
- Optimal bandwidth  $O(n^{-1/(2k+d)})$
- Optimal convergence rate  $O(n^{-k/(2k+d)})$

# Curse of dimensionality

Broadly, *two approaches* for handling the *curse of dimensionality*.

1. Enforce fairly strong *smoothness assumptions* on the model space (e.g., Hirano et al. 2003).
  - No general guarantee of *consistency*
2. Ensemble machine learning, e.g., *Super Learning* (van der Laan et al. 2007).
  - No guarantee of  $n^{-1/4}$  *convergence rates*

## An important class of functions

Consider space of *cadlag* functions with *finite variation norm*.

**Def.** *cadlag* = *left-hand continuous* with *right-hand limits*

**Variation norm** Let  $\theta_s(u) = \theta(u_s, 0_{s^c})$  be the *section* of  $\theta$  that sets the coordinates in  $s$  equal to zero.

The *variation norm* of  $\theta$  can be written:

$$|\theta|_v = \sum_{s \subset \{1, \dots, d\}} \int |d\theta_s(u_s)|,$$

where  $x_s = (x(j) : j \in s)$  and the sum is over all subsets.

## Variation norm

We can represent the function  $\theta$  as

$$\theta(x) = \theta(0) + \sum_{s \subset \{1, \dots, d\}} \int \mathbb{I}(x_s \geq u_s) d\theta_s(u_s),$$

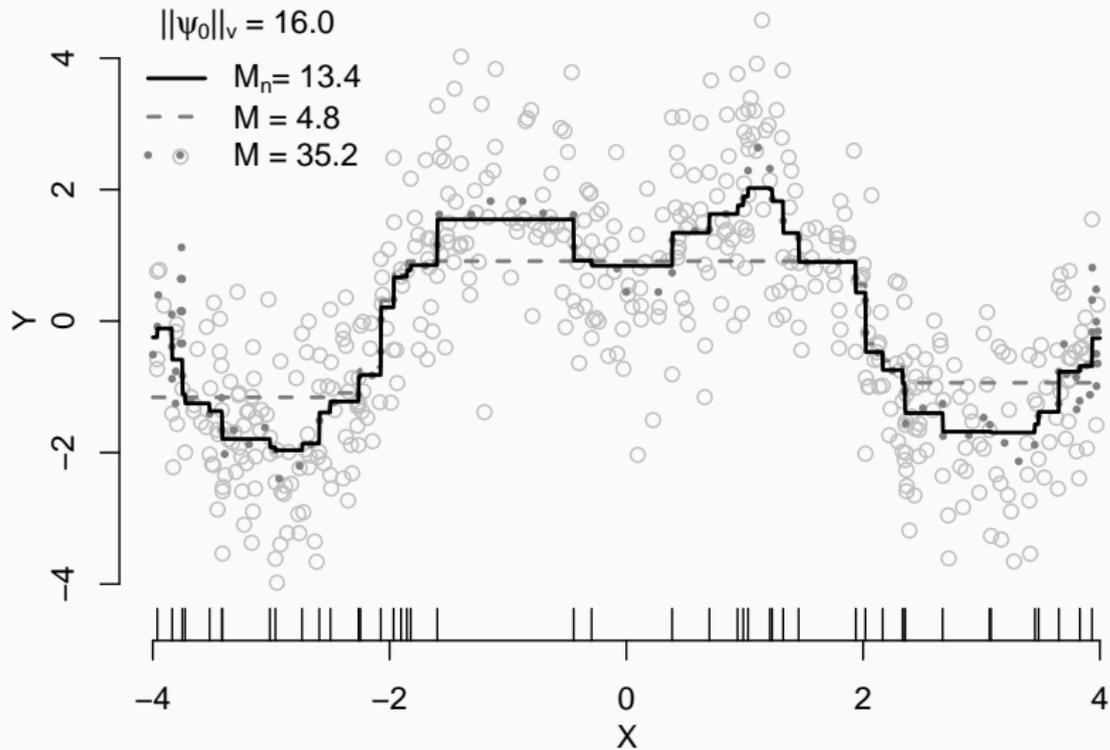
For discrete measures  $d\theta_s$  with *support points*  $\{u_{s,j} : j\}$  we get a *linear combination* of indicator *basis functions*:

$$\theta(x) = \theta(0) + \sum_{s \subset \{1, \dots, d\}} \sum_j \beta_{s,j} \theta_{u_{s,j}}(x),$$

where  $\beta_{s,j} = d\theta_s(u_{s,j})$ ,  $\theta_{u_{s,j}}(x) = \mathbb{I}(x_s \geq u_{s,j})$ , and

$$|\theta|_v = \theta(0) + \sum_{s \subset \{1, \dots, d\}} \sum_j |\beta_{s,j}|.$$

# HAL illustration



## Convergence rate of HAL

We have, for  $\alpha(d) = 1/(d + 1)$ ,

$$|\theta_{n,M} - \theta_{0,M}|_{P_0} = o_P(n^{-(1/4+\alpha(d)/8)}).$$

Thus, if we select  $M > |\theta_0|_V$ , then

$$|\theta_{n,M} - \theta_0|_{P_0} = o_P(n^{-(1/4+\alpha(d)/8)}) .$$

Due to oracle inequality for the cross-validation selector  $M_n$ ,

$$|\theta_{n,M_n} - \theta_0|_{P_0} = o_P(n^{-(1/4+\alpha(d)/8)}) .$$

Improved convergence rate (Bibaut and van der Laan 2019):

$$|\theta_{n,M_n} - \theta_0|_{P_0} = o_P(n^{-1/3} \log(n)^{d/2}) .$$

## HAL estimate of $g_{0,A}$

If the nuisance functional  $g_{0,A}$  is cadlag with finite sectional variation norm, logit  $g$  can be expressed (Gill et al. 1995):

$$\text{logit } g_\beta = \beta_0 + \sum_{s \subset \{1, \dots, d\}} \sum_{i=1}^n \beta_{s,i} \phi_{s,i},$$

where  $\phi_{s,i}$  is an indicator basis function.

The loss-based HAL estimator  $\beta_n$  may then be defined as

$$\beta_{n,\lambda} = \arg \min_{\beta: |\beta_0| + \sum_{s \subset \{1, \dots, d\}} \sum_{i=1}^n |\beta_{s,i}| < \lambda} P_n \mathcal{L}(\text{logit } g_\beta),$$

where  $\mathcal{L}(\cdot)$  is an appropriate loss function.

Denote by  $g_{n,\lambda} \equiv g_{\beta_{n,\lambda}}$  the HAL estimate of  $g_{0,A}$ .

## Targeted selection of $\lambda_n$ for IPW estimation

1. CV-based: choose  $\lambda_n$  as cross-validated empirical minimizer of negative log-density loss (Dudoit and van der Laan 2005):

$$\mathcal{L}(\cdot) = -\log(g_{n,A,\lambda}(A | W)).$$

n.b., “targeted” but incorrect tradeoff ( $g_{n,A,\lambda}$  instead of  $\psi_{n,\delta}$ ).

2. EIF-based: choose  $\lambda_n$  to solve the EIF estimating equation:

$$\lambda_n = \arg \min_{\lambda} |P_n D_{\text{CAR}}(g_{n,A,\lambda}, \bar{Q}_{n,Y})|,$$

where  $\bar{Q}_{n,Y}$  is an estimate of  $\bar{Q}_{0,Y}$  and  $D^* = D_{\text{IPW}} - D_{\text{CAR}}$ .

## Agnostic selection of $\lambda_n$ for IPW estimation

What if we dispensed with criteria based on  $\psi_{n,\delta}$  altogether?

1. Plateau-based: choose  $\lambda_n$  as the first in  $\lambda_1, \dots, \lambda_K$  s.t.

$$|\psi_{n,\lambda_{j+1}} - \psi_{n,\lambda_j}|_{j=1}^{K-1} \leq Z_{(1-\alpha/2)}[\sigma_{n,\lambda_{j+1}} - \sigma_{n,\lambda_j}]_{j=1}^{K-1},$$

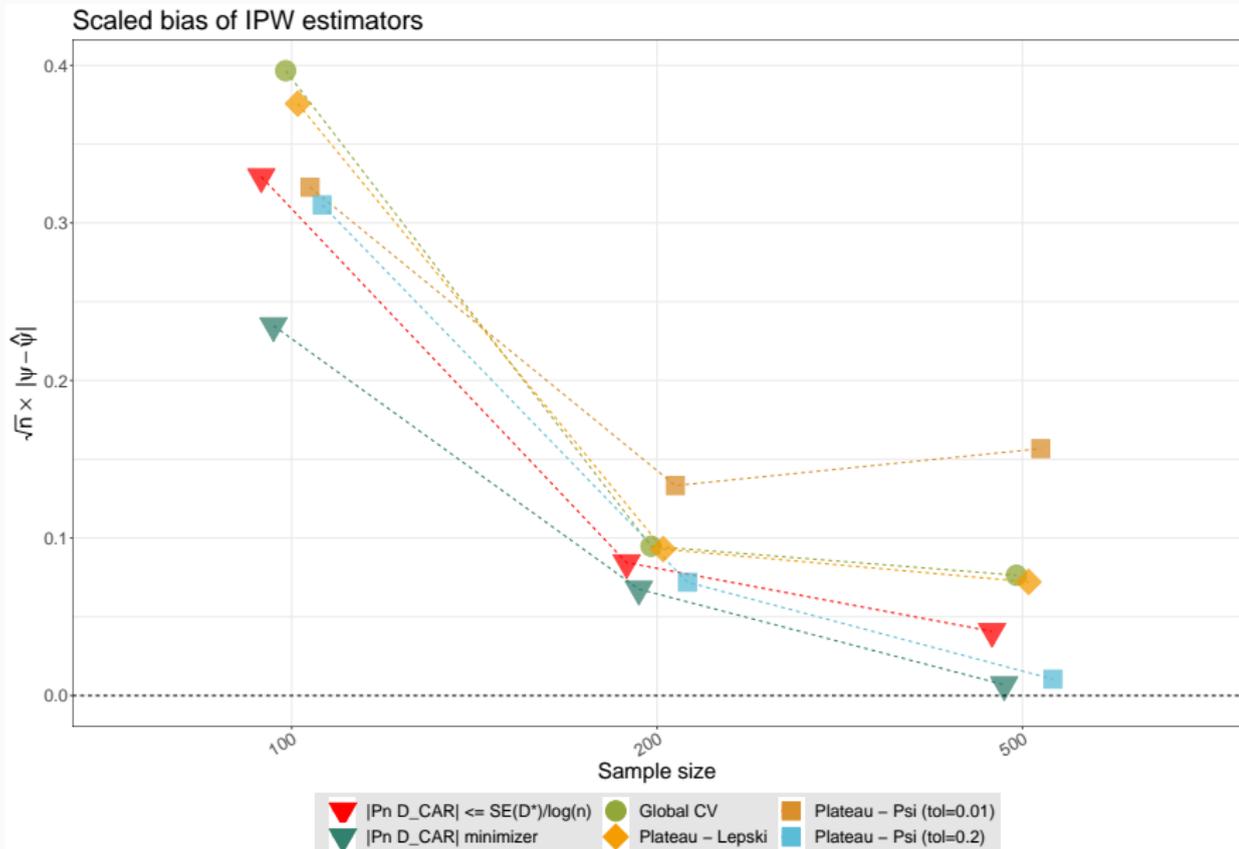
where  $\sigma_{n,\lambda_j}$  is a variance estimate at  $\lambda_j$ .

2. Plateau-based: choose  $\lambda_n$  as the first in  $\lambda_1, \dots, \lambda_K$  s.t.

$$\left[ \frac{|\psi_{n,\lambda_{j+1}} - \psi_{n,\lambda_j}|}{\max_j |\psi_{n,\lambda_{j+1}} - \psi_{n,\lambda_j}|} \right]_{j=1}^{K-1} \leq \tau$$

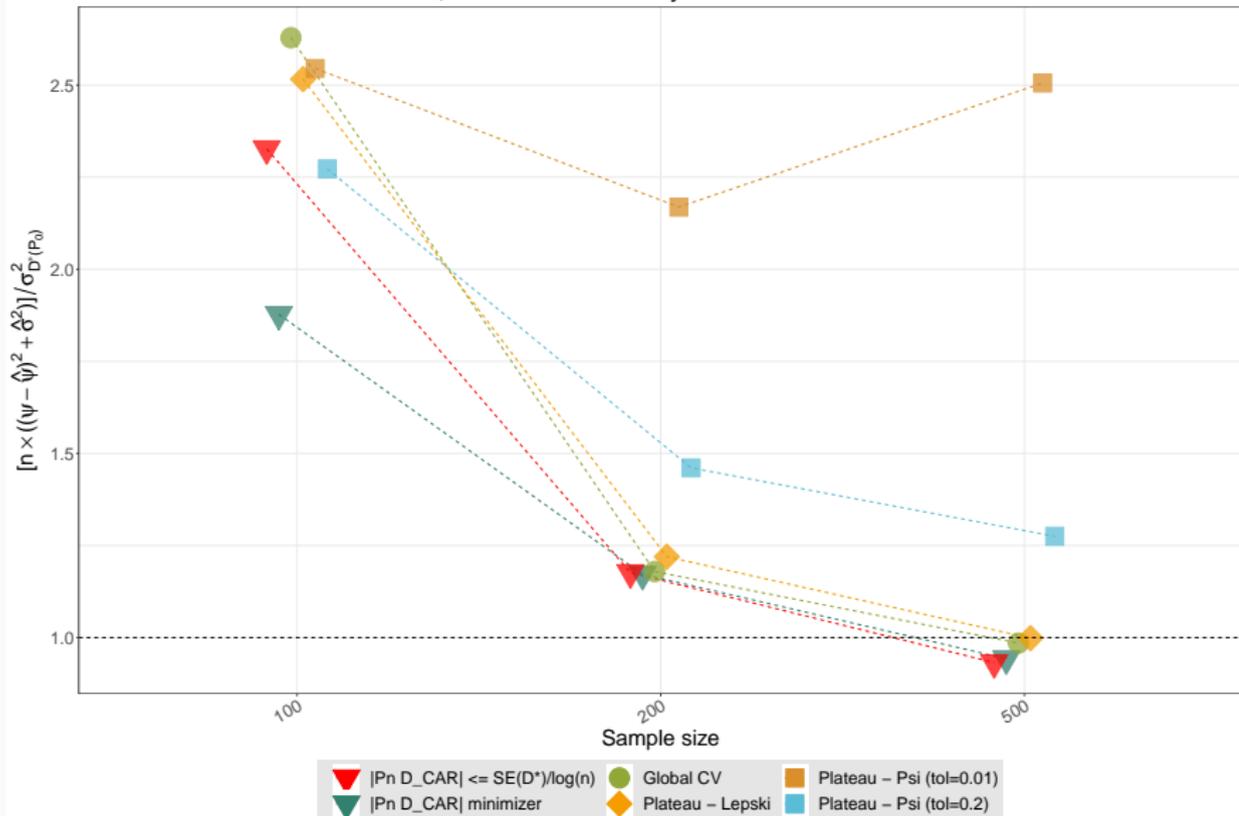
for an arbitrary tolerance level  $\tau$ .

# Simulation results: scaled bias



# Simulation results: relative MSE

Scaled MSE of IPW estimators, relative to efficiency bound



# The big picture

1. Unlike classical IPW estimators, ours avoid the asymptotic curse of dimensionality and are asymptotically efficient;
2. Our approach leverages flexible conditional density estimation for initial generalized propensity score estimates; and
3. In contrast with doubly robust estimators, our estimators can be formulated without the form of the EIF.
4. Check out the R packages that make this possible
  - `hal9001`: <https://github.com/tlverse/hal9001>
  - `haldensify`: <https://github.com/nhejazi/haldensify>

# Thank you!

 <https://nimahejazi.org>

 <https://twitter.com/nshejazi>

 <https://github.com/nhejazi>

 Manuscript coming soon — stay tuned!

# Appendix

# From the causal to the statistical target parameter

## **Assumption 1: *Stable Unit Treatment Value (SUTVA)***

- $Y_i^{\delta(a_i, w_i)}$  does not depend on  $\delta(a_j, w_j)$  for  $i = 1, \dots, n$  and  $j \neq i$ , or lack of interference (Rubin 1978; 1980)
- $Y_i^{\delta(a_i, w_i)} = Y_i$  in the event  $A_i = \delta(a_i, w_i)$ ,  $i = 1, \dots, n$

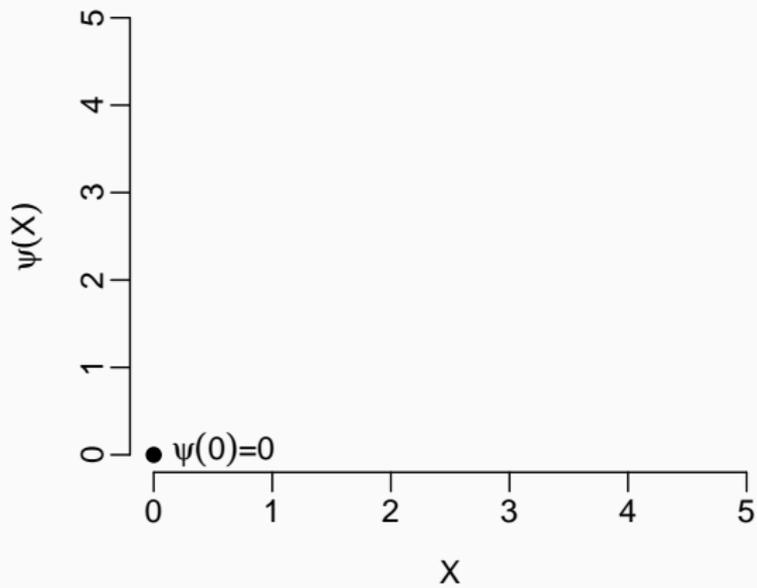
## **Assumption 2: *Ignorability***

$$A_i \perp\!\!\!\perp Y_i^{\delta(a_i, w_i)} \mid W_i, \text{ for } i = 1, \dots, n$$

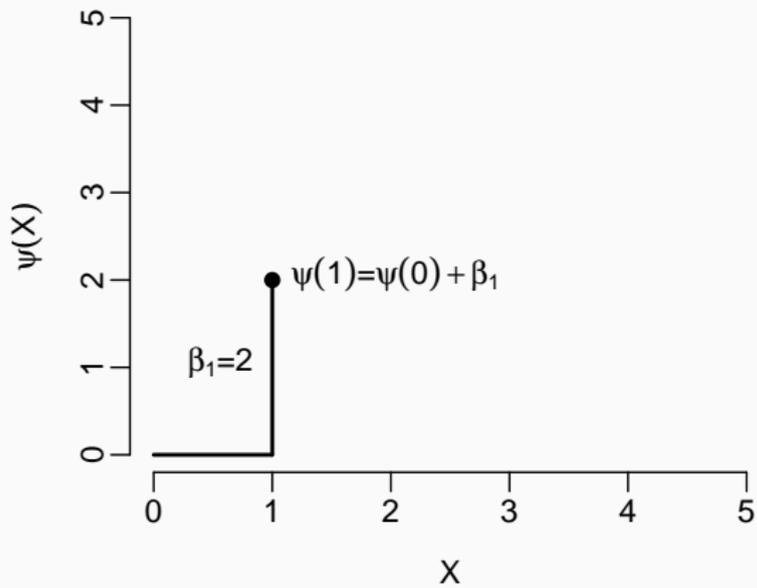
## **Assumption 3: *Positivity***

$a_i \in \mathcal{A} \implies \delta(a_i, w_i) \in \mathcal{A}$  for all  $w \in \mathcal{W}$ , where  $\mathcal{A}$  denotes the support of  $A$  conditional on  $W = w_i$  for all  $i = 1, \dots, n$

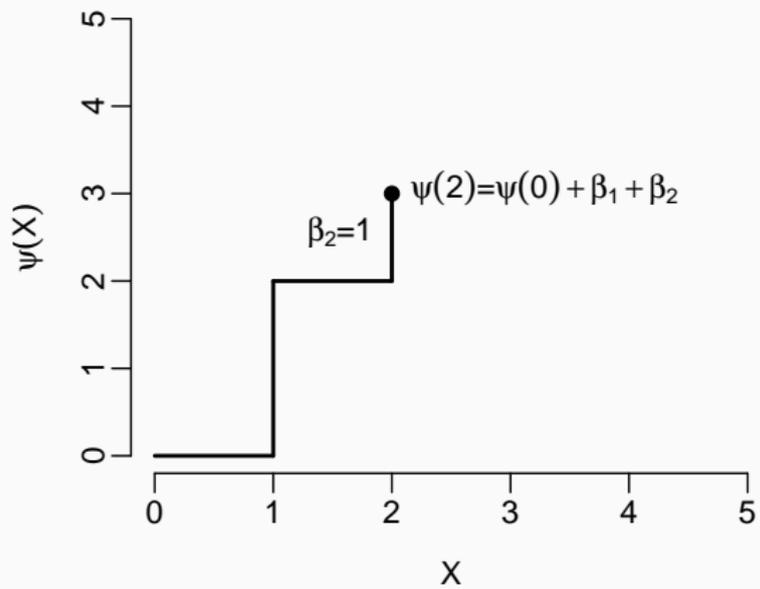
# HAL illustration



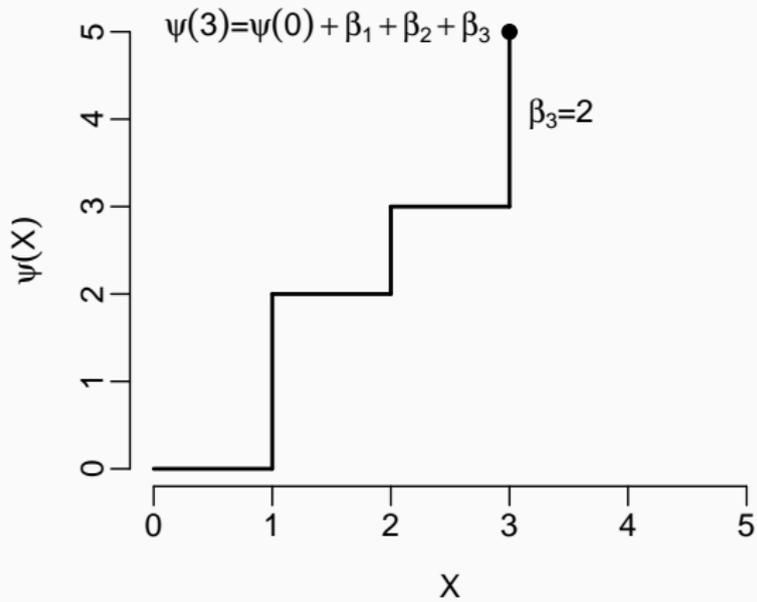
# HAL illustration



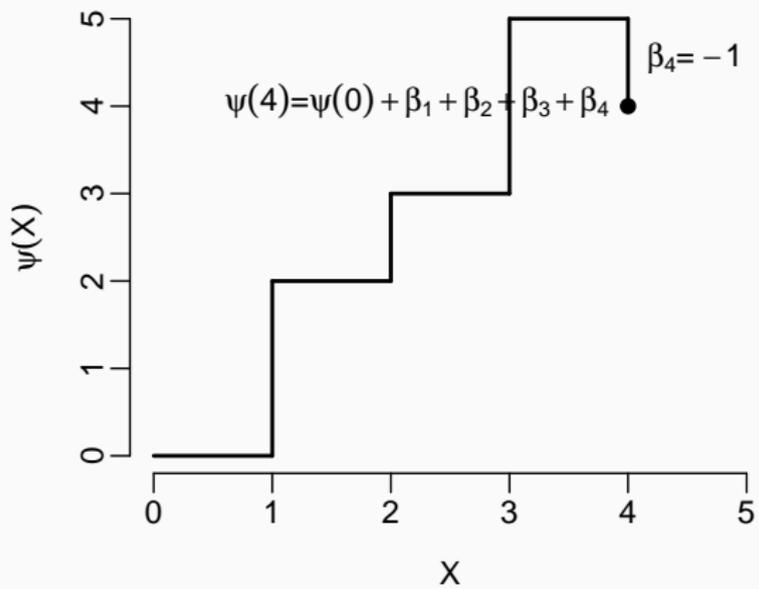
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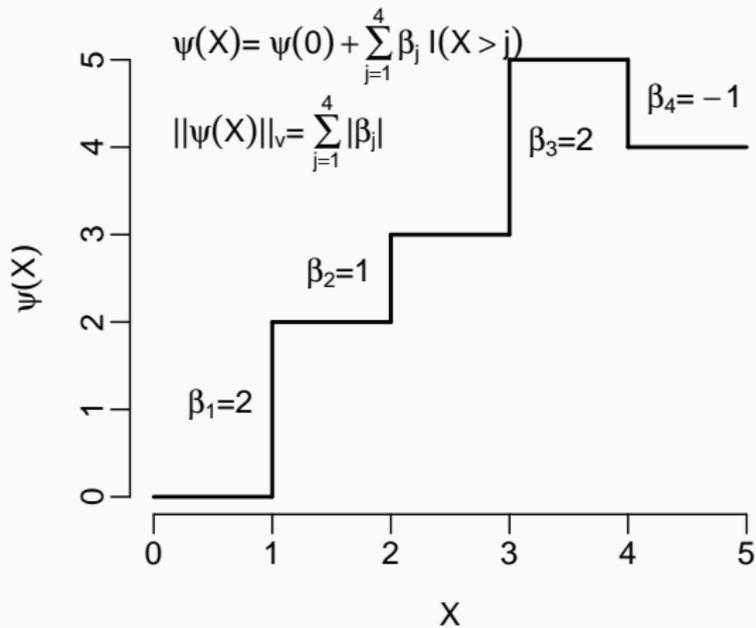
# HAL illustration



# HAL illustration



# HAL illustration



## Literature: Haneuse and Rotnitzky (2013)

- *Proposal*: Characterization of stochastic interventions as *modified treatment policies* (MTPs).
- Assumption of *piecewise smooth invertibility* allows for the intervention distribution of any MTP to be recovered:

$$g_{0,\delta}(a | w) = \sum_{j=1}^{J(w)} I_{\delta,j} \{h_j(a, w), I\} g_0 \{h_j(a, w) | I\} h'_j(a, w)$$

- Such intervention policies account for the natural value of the intervention  $A$  directly yet are interpretable as the imposition of an altered intervention mechanism.
- Identification conditions for assessing the parameter of interest under such interventions appear technically complex (at first).

## Literature: Young et al. (2014)

- Establishes equivalence between g-formula when proposed intervention depends on natural value and when it does not.
- This equivalence leads to a sufficient positivity condition for estimating the counterfactual mean under MTPs via the same statistical functional studied in Díaz and van der Laan (2012).
- Extends earlier identification results, providing a way to use the same statistical functional to assess  $\mathbb{E}Y_{\delta(A,W)}$  or  $\mathbb{E}Y_{\delta(W)}$ .
- The authors also consider limits on implementing shifts  $\delta(A, W)$ , and address working in a longitudinal setting.

## Literature: Díaz and van der Laan (2018)

- Builds on the original proposal, accomodating MTP-type shifts  $\delta(A, W)$  proposed after their earlier work.
- To protect against positivity violations, considers a specific shifting mechanism:

$$\delta(a, w) = \begin{cases} a + \delta, & a + \delta < u(w) \\ a, & \text{otherwise} \end{cases}$$

- Proposes an improved TMLE algorithm, with a single auxiliary covariate for constructing the TML estimator.

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